

From the INTERNATIONAL SEARCHING AUTHORITY STEPHANIE L. SEIDMAN, ESQ. FISH & RICHARDSON P.C. 12390 EL CAMINO REAL WRITTEN OPINION OF THE SAN DIEGO, CA 92130 INTERNATIONAL SEARCHING AUTHORITY (PCT Rule 43bis.1) Date of mailing (day/month/year) Applicant's or agent's file reference FOR FURTHER ACTION See paragraph 2 below 18202-022WO1 International application No. International filing date (day/month/year) Priority date (day/month/year) PCT/US04/22907 16 July 2004 (16.07.2004) 16 July 2003 (16.07.2003) International Patent Classification (IPC) or both national classification and IPC PC(7): A61K 31/47,31/385,31/35,31/075; C07D 311/04,335/04,215/00,215/16; C07C 41/00 and US Cl.: 514/434,456,311,717; 546/152,178; 549/23,434; 568/632 Applicant LIGAND PHARMACEUTICALS, INC 1. This opinion contains indications relating to the following items: Box No. I Basis of the opinion Box No. II Box No. III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability Box No. IV Lack of unity of invention Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial Box No. V applicability; citations and explanations supporting such statement Box No. VI Certain documents cited Box No. VII Certain defects in the international application Box No. VIII Certain observations on the international application 2. FURTHER ACTION If a demand for international preliminary examination is made, this opinion will be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA") except that this does not apply where the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notified the International Bureau under Rule 66.1bis(b) that written opinions of this International Searching Authority will not be so considered. If this opinion is, as provided above, considered to be a written opinion of the IPEA, the applicant is invited to submit to the IPEA a written reply together, where appropriate, with amendments, before the expiration of 3 months from the date of mailing of Form PCT/ISA/220 or before the expiration of 22 months from the priority date, whichever expires later. For further options, see Form PCT/ISA/220. 3. For further details, see notes to Form PCT/ISA/220.

Authorized officer

Telephone No.

Form PCT/ISA/237 (cover sheet) (January 2004)

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Commissioner for Patents

P.O. Box 1450



WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/US04/22907

Box N	. I Basis of this opinion	
1. With was f	egard to the language, this opinion has been established on the basis of the international application in the language in which it led, unless otherwise indicated under this item.	ţ
	This opinion has been established on the basis of a translation from the original language into the following language which is the language of a translation furnished for the purposes of international search (under Rules 12.3 and 23.1(b)).	
2. With inven	egard to any nucleotide and/or amino acid sequence disclosed in the international application and necessary to the claimed ion, this opinion has been established on the basis of:	
a.	type of material	
	a sequence listing	
	table(s) related to the sequence listing	
ъ.	format of material	
	in written format	
	in computer readable form	
c.	time of filing/furnishing	
	contained in international application as filed.	
	filed together with the international application in computer readable form.	
	furnished subsequently to this Authority for the purposes of search.	
3.	In addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto has been filed or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished.	
4. Additi	onal comments:	
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Box No. V Reasoned statement under Rule 43 bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement 1. Statement Novelty (N) Claims 1,8,15,23,30,38 YES Claims 2-7,9-14,16-22,24-29,31-37,39-114 Inventive step (IS) Claims 1.8.15,23,30,38 YES Claims 2-7.9-14.16-22.24-29.31-37.39-114 NO Industrial applicability (IA) Claims 1-114 YES Claims NONE NO

2. Citations and explanations:

Claims 1,8,15,23,30,38 lack novelty under PCT Article 33(2) as being anticipated by Chandraratna USP 5,015,658. the compounds are pharmaceuticals.

Claims 1,8,15,23,30,38 lack novelty under PCT Article 33(2) as being anticipated by Baldo et al USP 6,391,287 B1. A few of the species are listed below, although the reference teaches several. The compounds are cosmetic compositions.

Claims 1-114 meet the criteria set out in PCT Article 33(4), and thus have industrial applicability because the subject matter claimed can be made or used in industry.

Form PCT/ISA/237 (Box No. V) (January 2004)